

CGGGCTTATT	ATTCCACTGA	TGAGAACCTG	ATCCTTTCCC	CACTCCTGGG	TAACGTCTGC
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GACACCTTTG	GTGACATTAA	TTACCAAGAA	TTTGCTAAAA	GACTCTGGGG	TGACATCTAC
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GACACCAGCC	TCCCACGGAC	CCTAGACGAG	CTTGGCATCC	ACCTGACGAA	GGAGGAGCTG
AAGCTGAACA	TCCGCCCCTT	GCTCAGGCTG	GTCTGCAAAA	AGTTCTTTGG	CGAGTTCACA
GGCTTTGTGG	ACATGTGTGT	GCAGCATATC	CCTTCTCCAA	AGGTGGGCGC	CAAGCCCCAAG
ATTGAGCACA	CCTACACCGG	TGGTGTGGAC	TCCGACCTCG	GCGAAGCTAT	GAGTGACTGT
GACCCTGATG	GCCCCCTGAT	GTGCCACACT	ACTAAGATGT	TCAGCACACA	TGATGGAGTC
CAGTTTCACC	CCTTTGGCCG	GGTGCTGAGT	GGCACCATTG	ATGCTGGGCA	GCCTGTGAAG
GTTCTGGGGG	AGAACTACAC	CCTGGAGGAT	GAGGAAGACT	CCCCAATTTG	CCCCGTGGGC
CGCCTTTGGA	TCTCTGTGGC	CAGCTACCAC	ATCGAGGTGA	ACCGTGTTCC	TGCTGGCAAC
TGGGTTCTGA	TTGAAGGTGT	TGATCAACCA	ATTGTGAAGA	CAGCAACCAT	AACCGAACCC
CGAGGCAATG	AGGAGGCTCA	GATTTTCCGA	CCCTTGAAGT	TCAATACCAC	ATCTGTTATC
AAGATTGCTG	TGGAGCCAGT	CAACCCCTCA	GAGCTGCCCA	AGATGCTTGA	TGGCCTGCGC
AAGGTCAACA	AGAGCTATCC	ATCCCTCACC	ACCAAGGTGG	AGGAGTCTGG	CGAGCATGTG
ATCCTGGGCA	CTGGGGAGCT	CTACCTGGAC	TGTGTGATGC	ATGATTTGCG	GAAGATGTAC
TCAGAGATAG	ACATCAAGGT	GGCTGACCCA	GTTGTCACGT	TTTGTGAGAC	GGTCGTGGAA
ACATCCTCCC	TCAAGTGCTT	TGCTGAAACG	CCTAATAAGA	AGAACAAGAT	CACCATGATT
GCTGAGCCTC	TTGAGAAGGG	CCTGGCAGAG	GACATAGAGA	ATGAGGTGGT	CCAGATTACG
TGGAACAGGA	AGAAGCTGGG	AGAGTTCTTC	CAGACCAAGT	ACGATTGGGA	TCTGCTGGCT
GCCCGTTCCA	TCTGGGCTTT	TGGCCCTGAT	GCGACTGGCC	CCAACATTCT	GGTGGATGAT
ACTCTGCCCT	CTGAGGTGGA	CAAGGCTCTT	CTTGGTTCAG	TGAAGGACAG	CATCGTTCAA
GGT					

Fig 1A

TTGGGGACAT	TTTGGGGTGA	CACACTGAAC	TGCTGGATGC	TATCAGCATT	TAGTAGGTAT
GCTCGATGTC	TTGCAGAAGG	ACATGATGGT	CCTACACAGT	AAGGAATGGA	TTACCTACAA
TATTAATAGC	AGCCTCCCAT	ACACACTTTT	GACACCCTTC	CCTAAAGGAT	TAATATGCTC
CAACCTTCCT	GTCCCCACAG	TTCAGTGGCT	CTCCCTACCC	TCACCATGAT	CGGATGAAAA
AAAATAAGGT	TTCACAGCTT	AAGAGTGAAA	TTCTGGAATC	CAACTACAAG	CTCATAACTG
TAGCATGGAA	CCTGGTAGTA	GCATAATAAA	TAAATTTTTA	GTAAGAGGCT	TAAGAAATTT
TAGCAAAAAA	AGCACTCCCT	TTCTTCCTCC	CTACATATCT	CATATGTTTT	TCAACACAAA
AAATTCTGTG	ATTTTAGAGA	AACTTCTTAC	AGTACTTTTA	AGTTCAAAAC	CAGATGCTCA
TTACAGTTCT	TTTAAACACC	AAACTAGTCA	TCTCAAAAAT	ATGGCTAACT	CTCTGGACTA
AATTCCATAG	GAAAAATTAT	TAATTTCAAA	ATGCCTAATT	TTTGATCAAT	GCTGAAGAGC
CAAGCAATCA	TGTCCTGCTT	CTCACTCAGG	GCAGAGTTCT	CAGGTCAGAA	GCTCCGGAGT
CTGTCAGAGA	TTAAAATATC	ATCTCAACAA	TTCACAAGCT	ACTTCTAAGT	GTTACCCTAA
ATTAGTCACT	AATCGTTTCT	CCCCCAACTC	TATTTACAAA	ATTAAAGTTT	ACAGAATTGA
CAAAAACCAA	ACCAATGAAA	CAACCCAGGC	TATTTGCAGG	GGGGGGGAAA	GAGATACCCC
AAAAGTCAAC	CCTATTTACA	CGTAGTTAAA	AGAGTGATCC	AACAGATATT	ACCCTCCATA
AAGTACCTAA	AGGCAGGAGC	CGGAATT			

Fig. 1B

TTGGGGACAT TTTGGGGTGA CACACTGAAC TGCTGGATGC TATCAGCATT TAGTAGGTAT
GCTCGATGTC TTGCAGAAGG ACATGATGGT CCTACACAG

Fig. 1C

TTGGGGACAT TTTGGGGTGA CACACTGAAC TGCTGGATGC TATCAGCATT TAGTAGGTAT
GCTCGATGTC TTGCAGAAGG ACATGATGGT CCTACACAG

R A Y Y S T D E N L I L S
 P L L G N V C F S S S Q Y
 S I C F T L G S F A K I Y
 A D T F G D I N Y Q E F A
 K R L W G D I Y F N P K T
 R K F T K K A P T S S S Q
 R S F V E F I L E P L Y K
 I L A Q V V G D V D T S L
 P R T L D E L G I H L T K
 E E L K L N I R P L L R L
 V C K K F F G E F T G F V
 D M C V Q H I P S P K V G
 A K P K I E H T Y T G G V
 D S D L G E A M S D C D P
 D G P L M C H T T K M Y S
 T H D G V Q F H P F G R V
 L S G T I H A G Q P V K V
 L G E N Y T L E D E E D S
 P I C P V G R L W I S V A
 R Y H I E V N R V P A G N
 W V L I E G V D Q P I V K
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 Q I F R P L K F N T T S V
 I K I A V E P V N P S E L
 P K M L D G L R K V N K S
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 H V I L G T G E L Y L D C
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 P N K K N K I T M I A E P
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 V Q I T W N R K K L G E F
 F Q T K Y D W D L L A A R
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 A L L G S V K D S I V Q G

Fig. 2 A

Fig. 3A

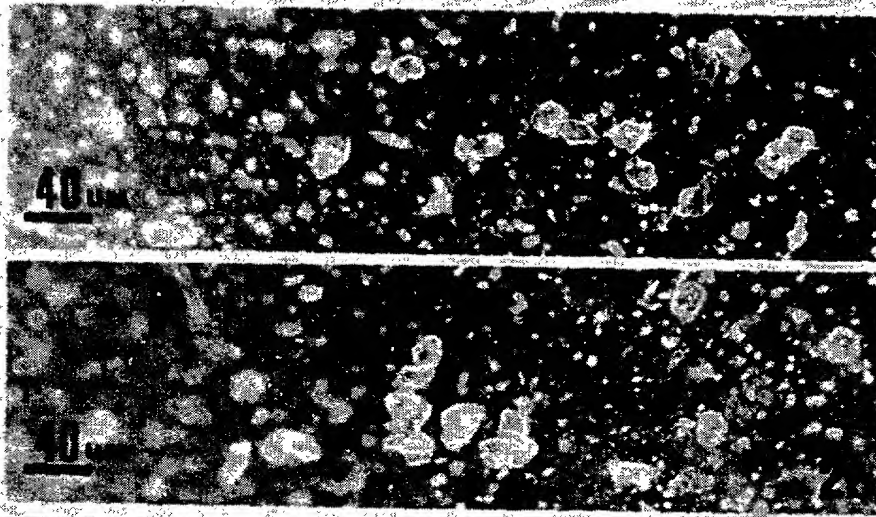
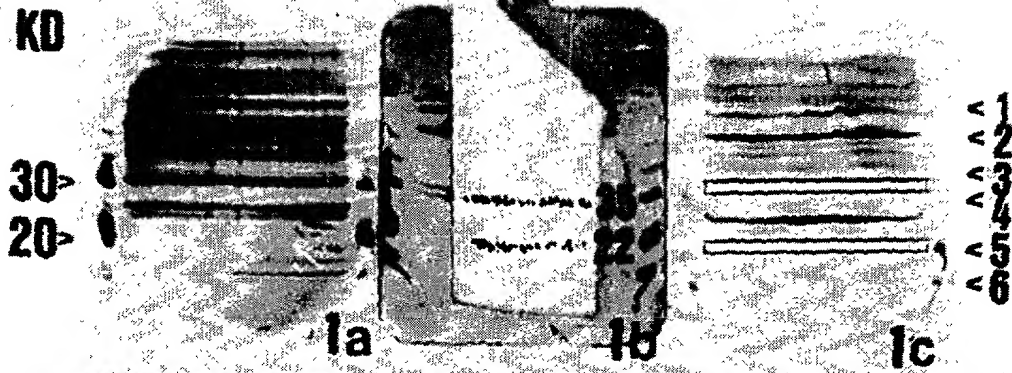


Fig. 3B

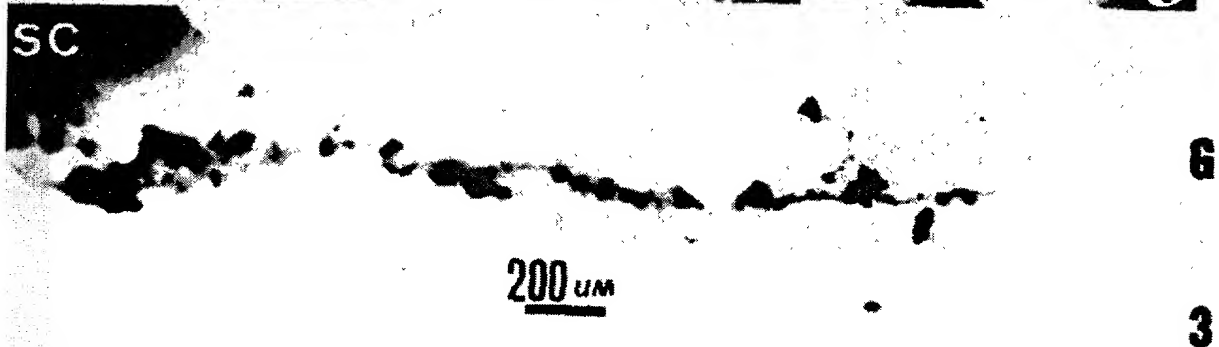
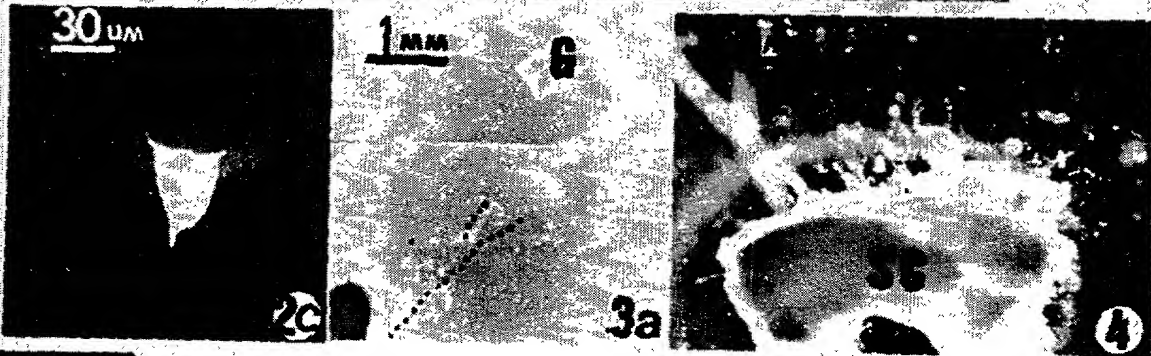


Fig. 3C



Fig. 4

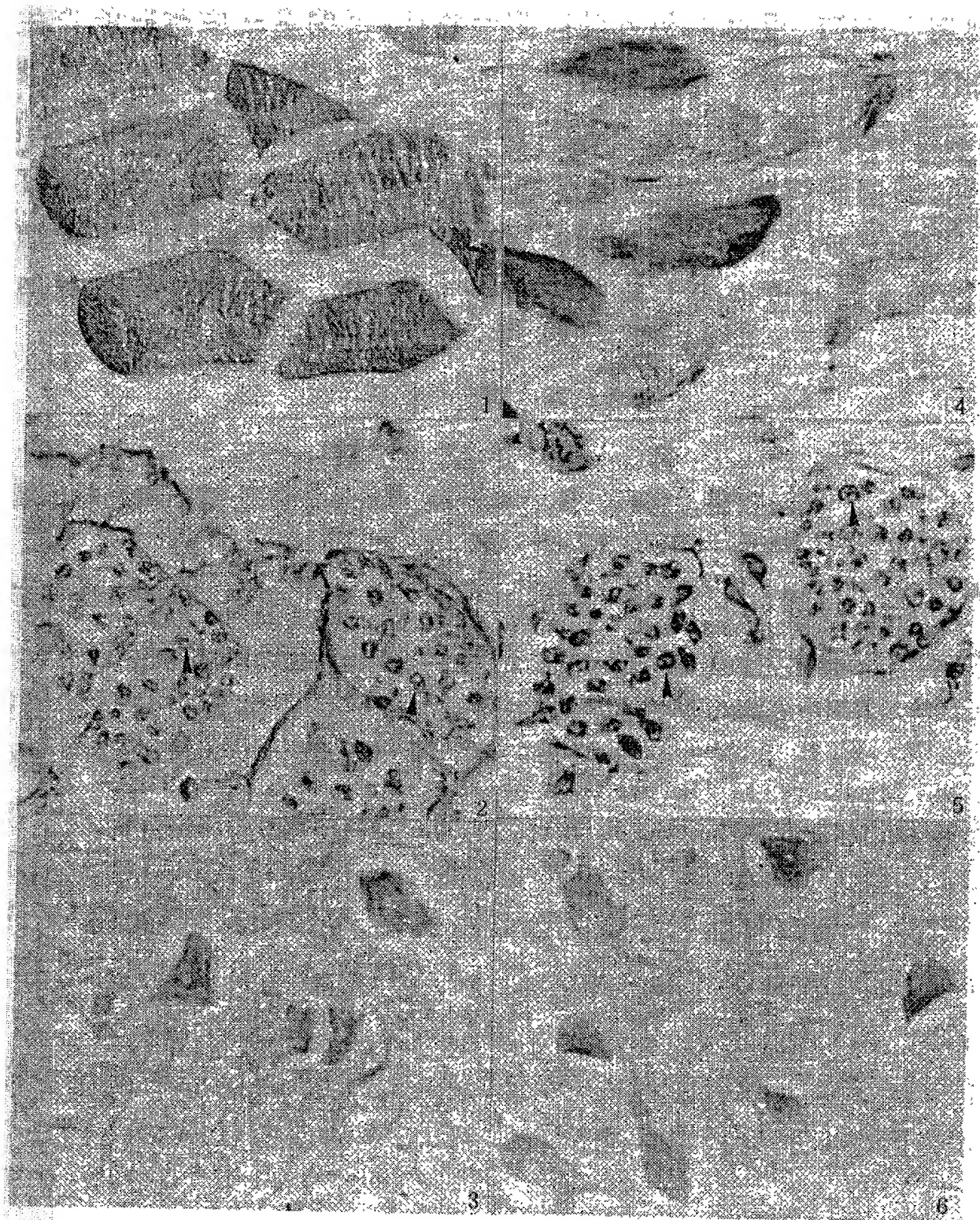


Fig. 5



Fig. 6

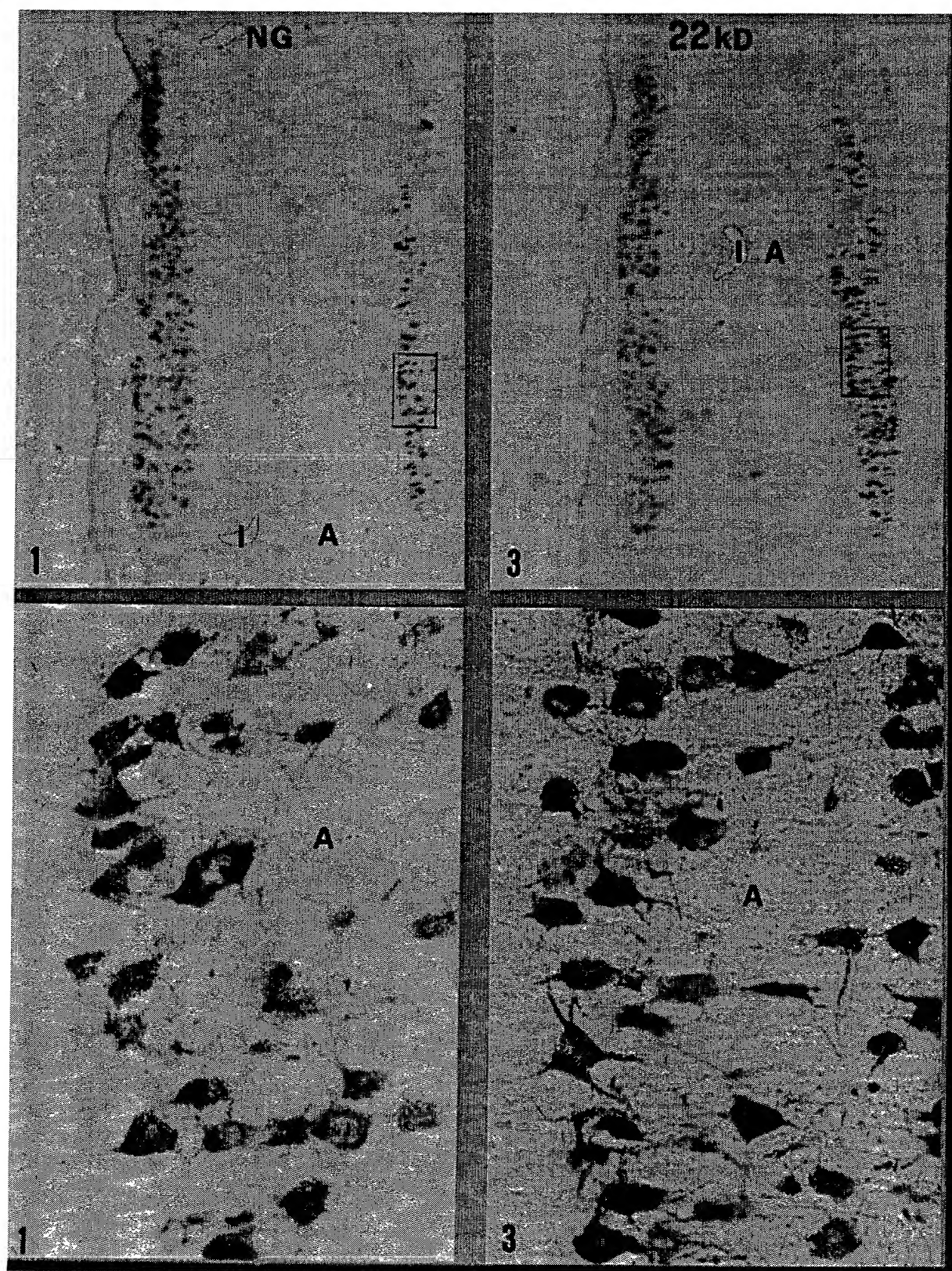


Fig. 7A

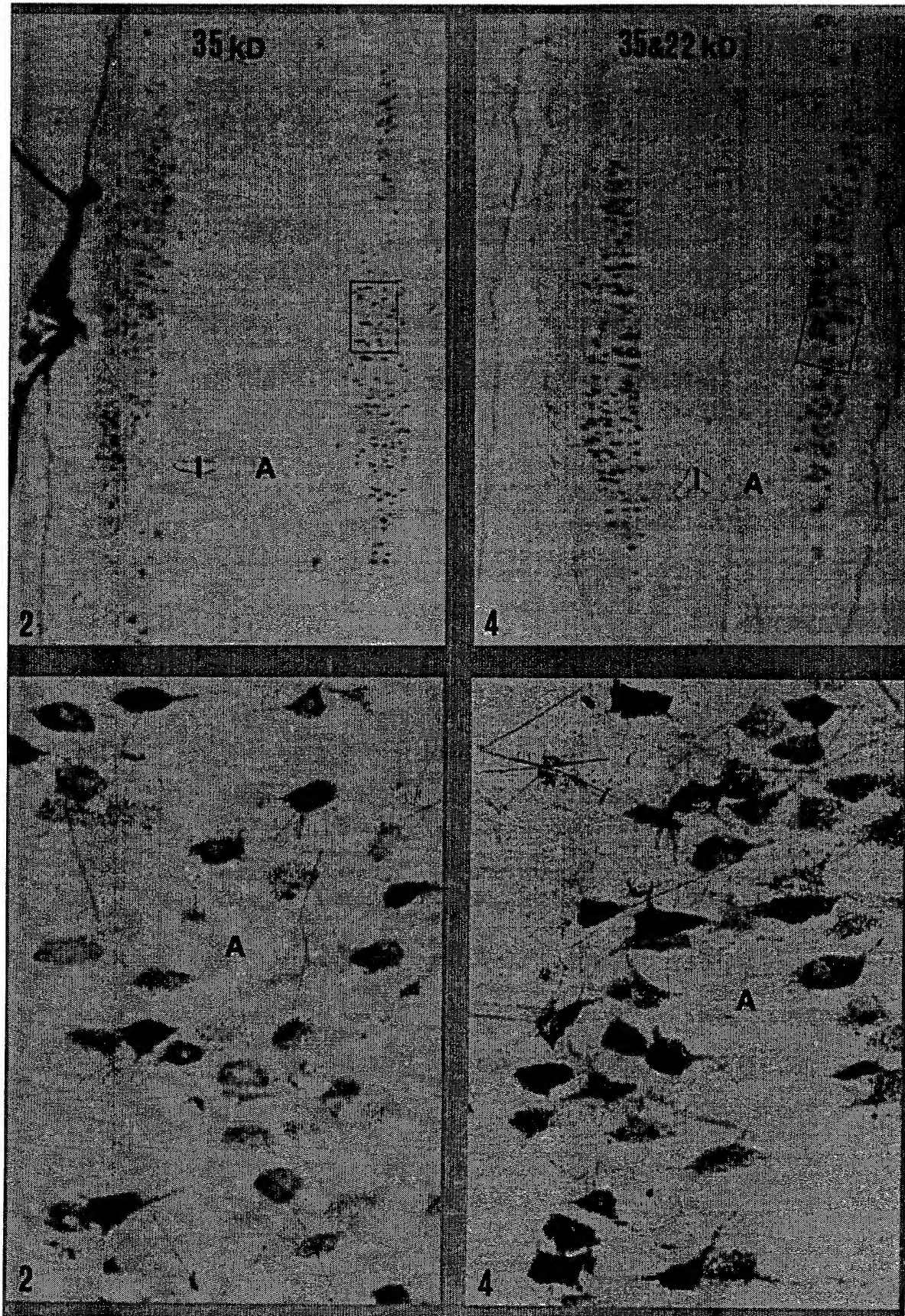


Fig. 7B

GROUP	NUMBER	% SURVIVAL
CONTROL	7	44.6 ± 7.5 (a)
35kD (MNTF1)	7	76.5 ± 10.8
22kD (MNTF2)	7	71.3 ± 8.7
35kD + 22kD	7	86.8 ± 5.8

SURVIVAL VALUES ARE MEAN ± S.D.

p<0.01

Fig. 8A

GROUP	% SURVIVAL	SD
CONTROL	44.6	7.5
35kD (MNTF1)	76.5	10.8
22Kd (MNTF2)	71.3	8.7
35kD + 22kD*	86.8	5.8

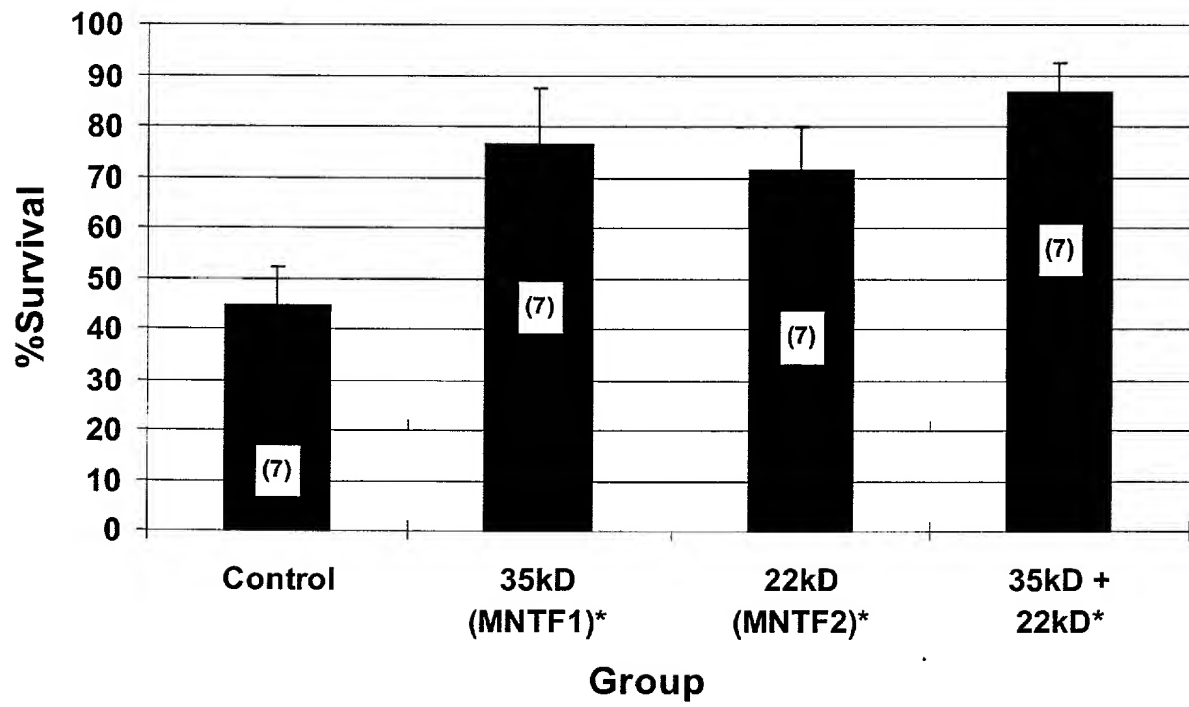


Fig. 8B

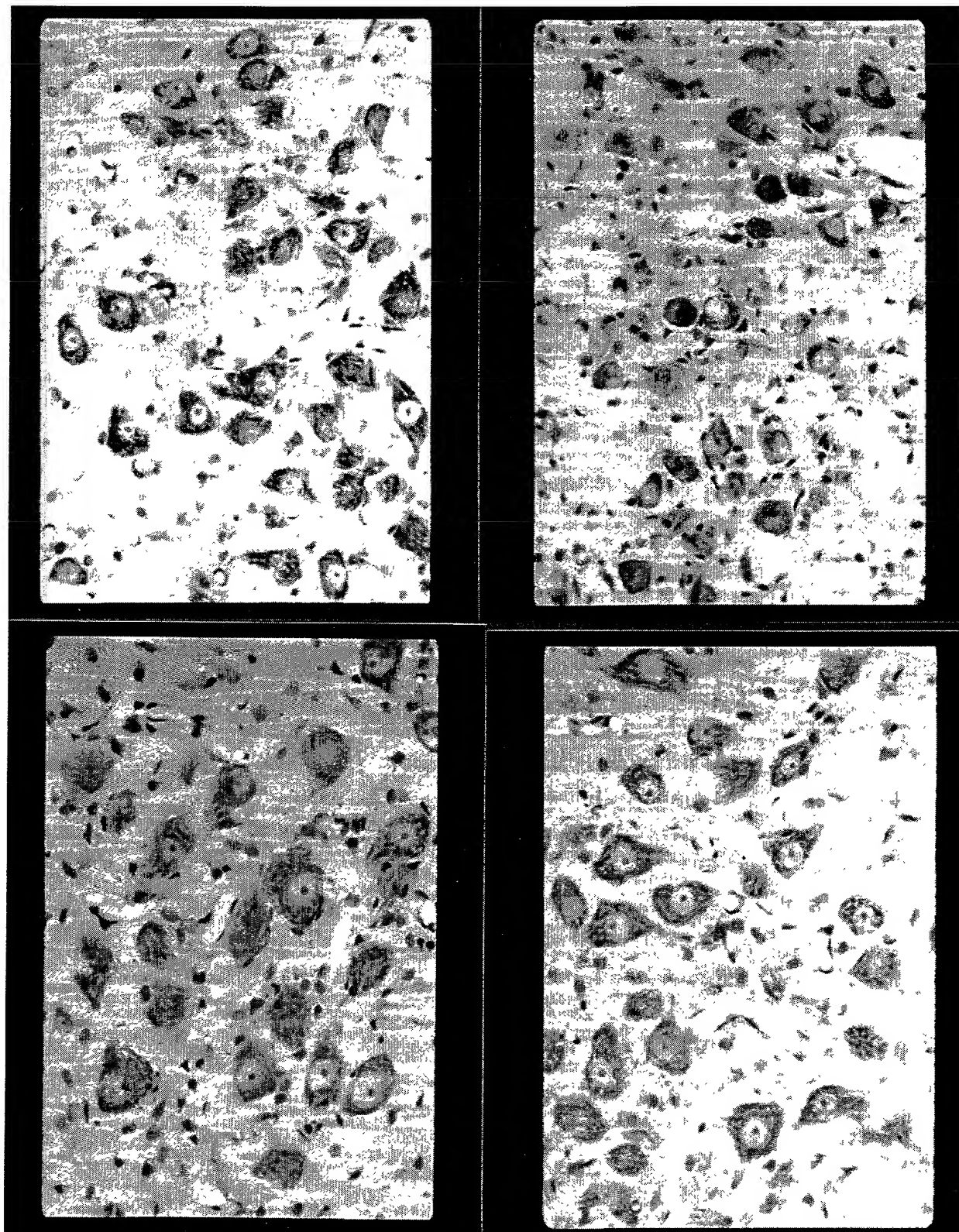


Fig. 9

% MOTONEURON SURVIVAL

TREATMENT	1 WEEK PO (n)	2 WEEKS PO (n)
CONTROL	53.4 ± 13.2 (3)	35.3 ± 9.2 (a)
35kD	70.9 ± 10.4 (4)	57.1 ± 6.3 (4) ^a
22kD	67.6 ± 10.4 (5)	56.3 ± 4.3 (3) ^a
35kD + 22kD	64.2 ± 6.6 (5)	53.1 ± 0.4 (3) ^a

VALUES ARE MEAN ± S.D.

p<0.01

Fig. 10A

TREATMENT	1 WEEK PO	2 WEEKS PO	SD (1 wk PO)	SD (2 wks PO)
CONTROL	53.4	35.3	13.2	9.2
35kD (MNTF1)	70.9	57.1	10.4	6.3
22kD (MNTF2)	67.6	56.3	10.4	4.3
35kD + 22kD*	64.2	53.1	6.6	0.4

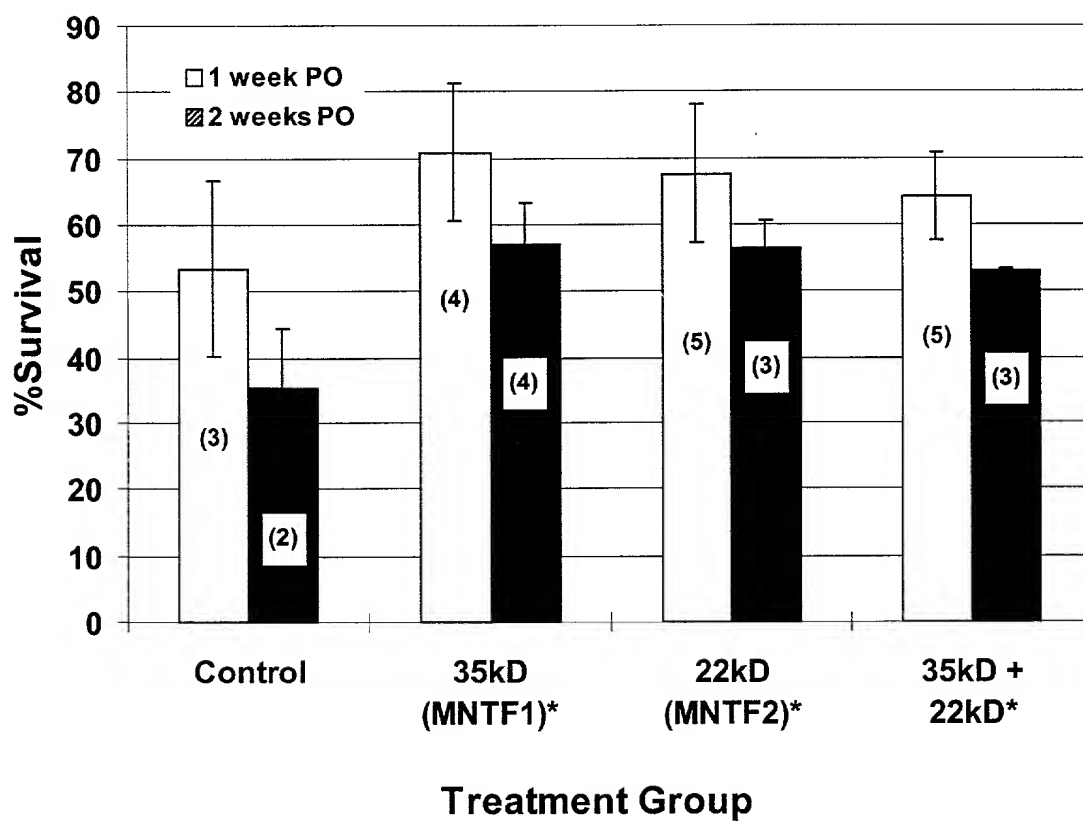


Fig. 10B

TREATMENT	1 WEEK PO	2 WEEKS PO	SD (1 wk PO)	SD (2 wks PO)
CONTROL	53.4	35.3	13.2	9.2
35kD (MNTF1)	70.9	57.1	10.4	6.3
22kD (MNTF2)	67.6	56.3	10.4	4.3
35kD + 22kD*	64.2	53.1	6.6	0.4

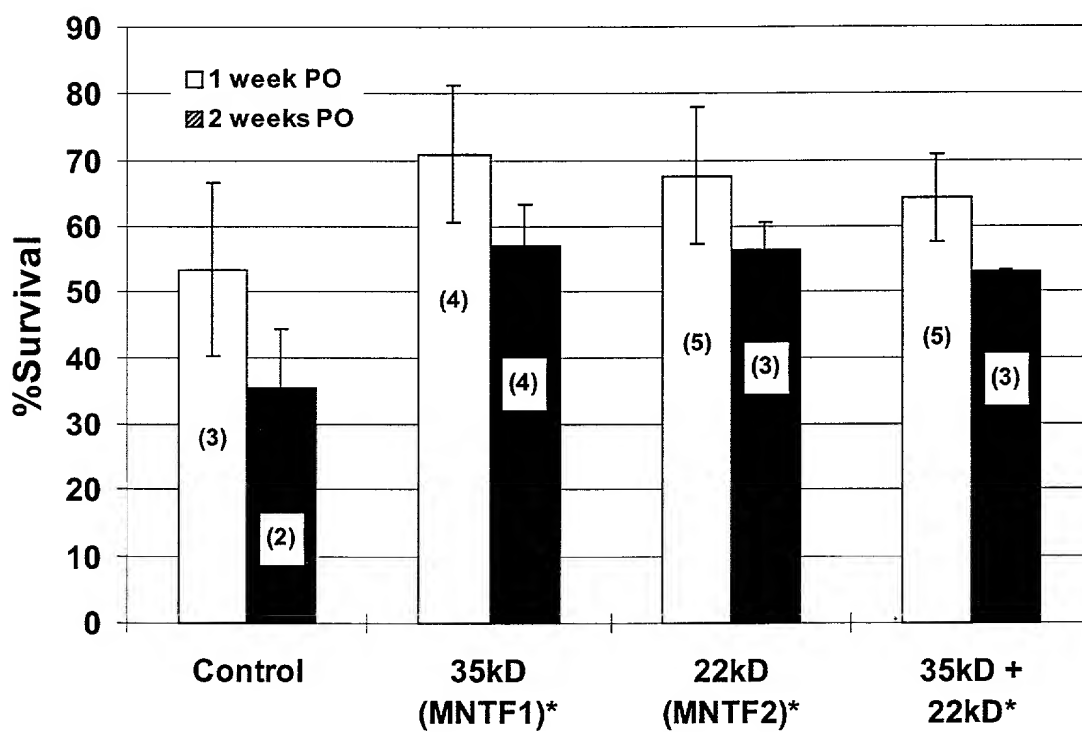


Fig. 10B

% MOTONEURON SURVIVAL

TREATMENT	2 WEEK PO	(n)
35kD + GOAT ANTI-RABBIT IgG	80.1 ± 11.3	(5) ^b
22kD + ANTI-35kD Mab	49.4 ± 0.9	(5) ^b

VALUES ARE MEAN ± S.D.

p<0.001

ANTI-35kD Mab = MONOCLONAL ANTIBODY DIRECTED
AGAINST 35 kD MOTONEURONOTROPHIC FACTOR.

Fig. 11A

TREATMENT	2 WEEKS PO	SD
35kD + GOAT ANTI-RABBIT IgG*	80.1	11.3
22kD + ANTI-35kD Mab*	49.4	0.9

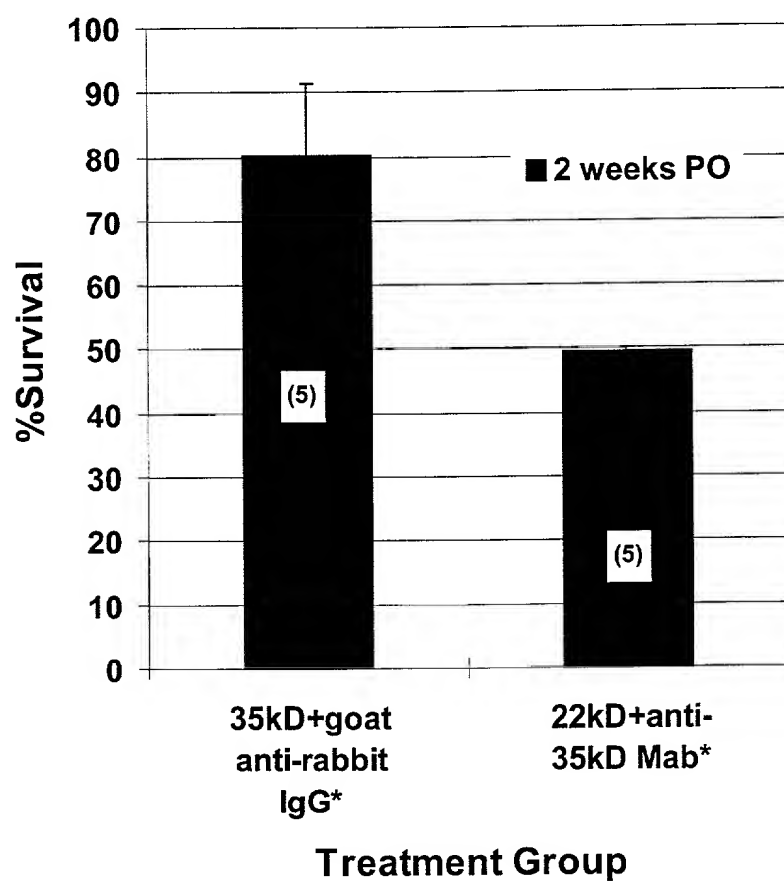


Fig. 11B

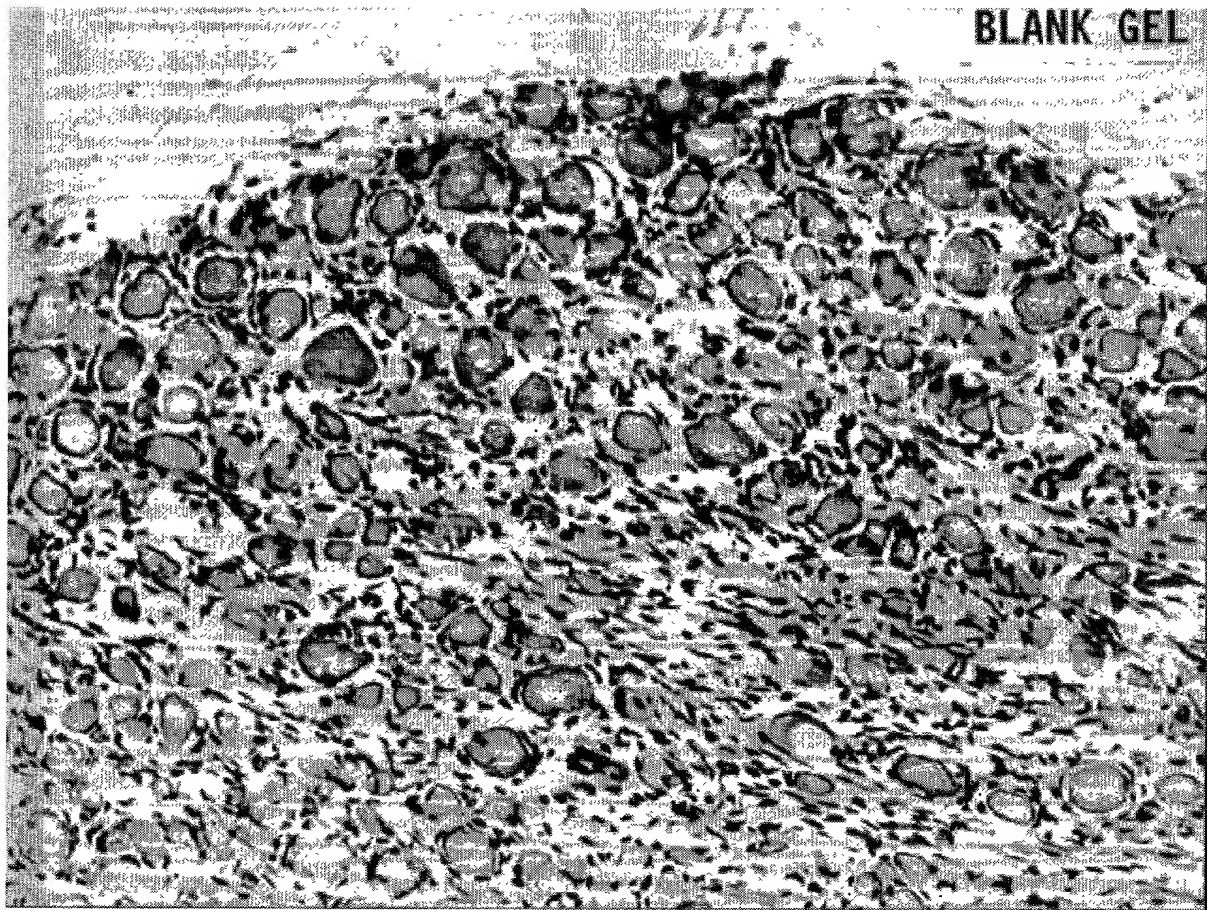


Fig. 12A

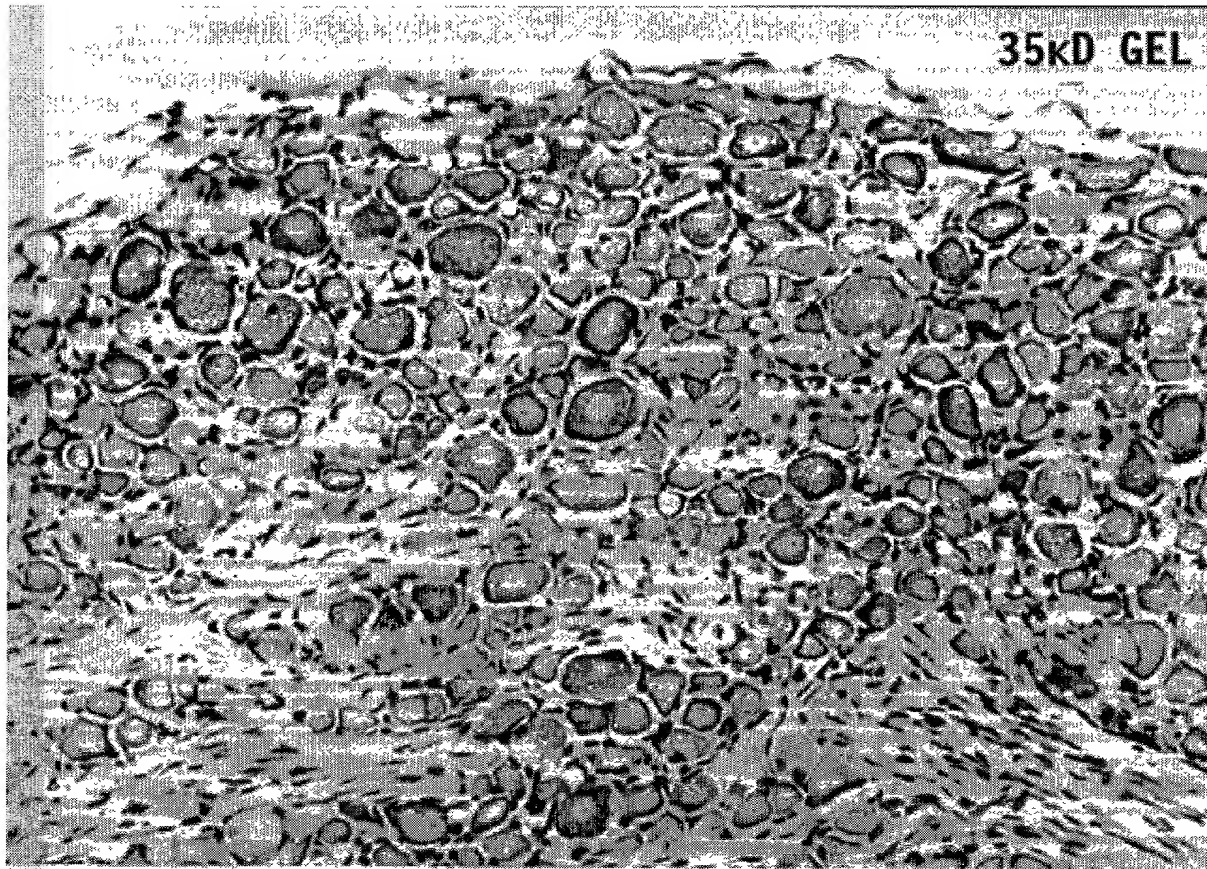


Fig. 12B

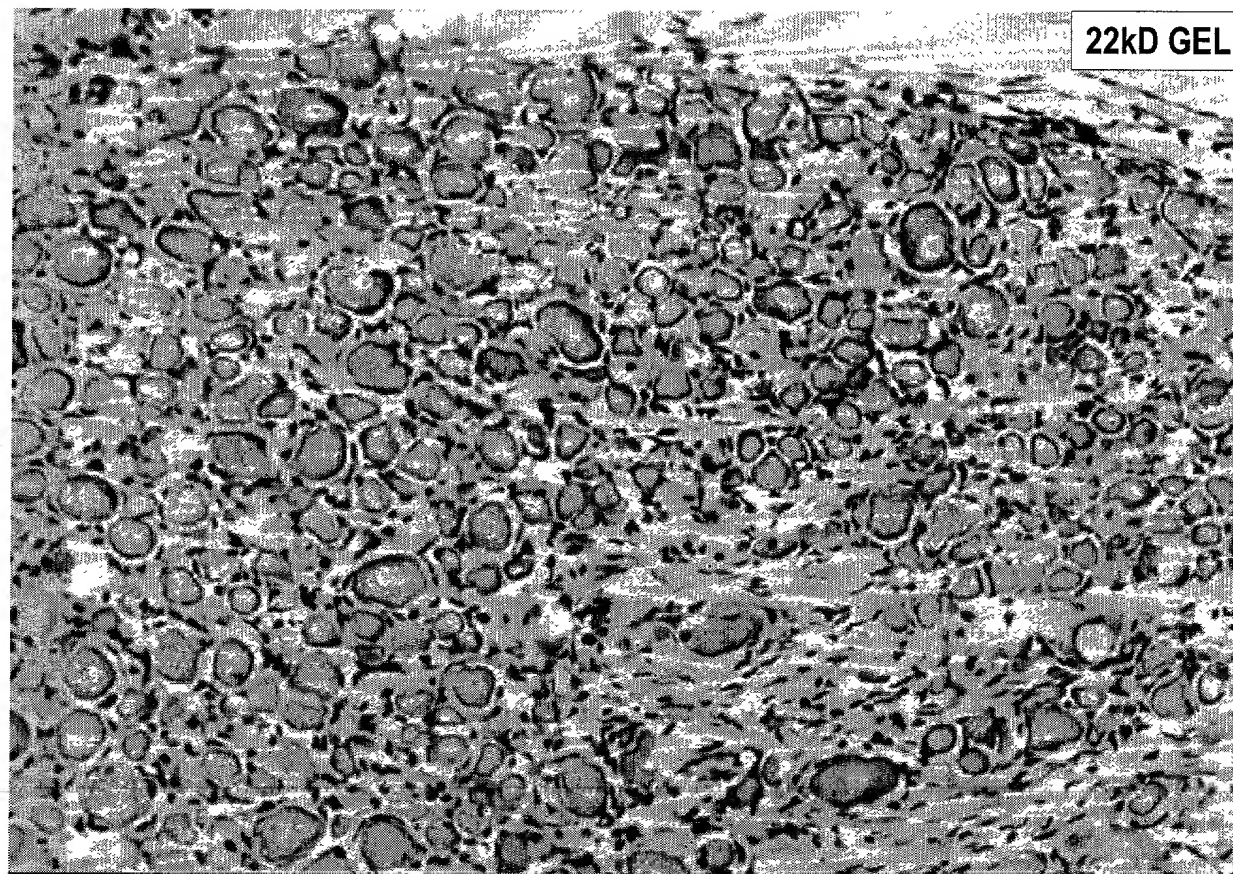


Fig. 12C

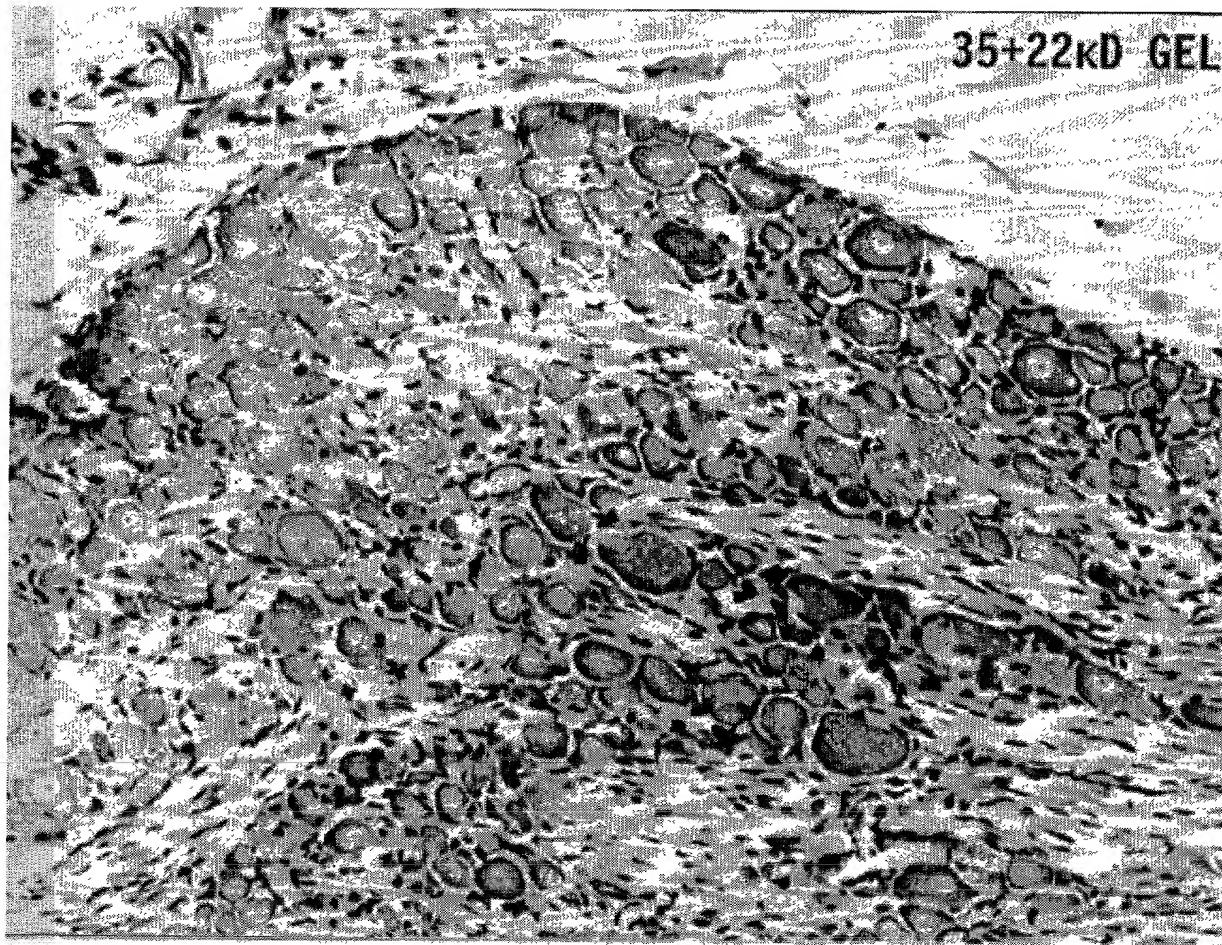


Fig. 12D

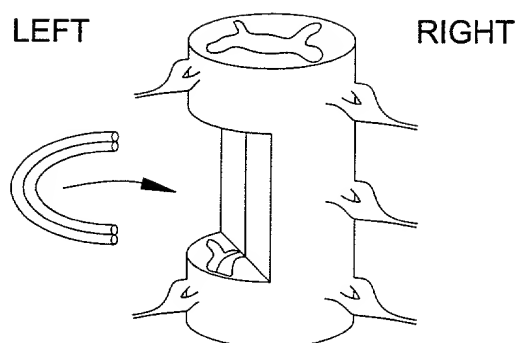


Fig. 13

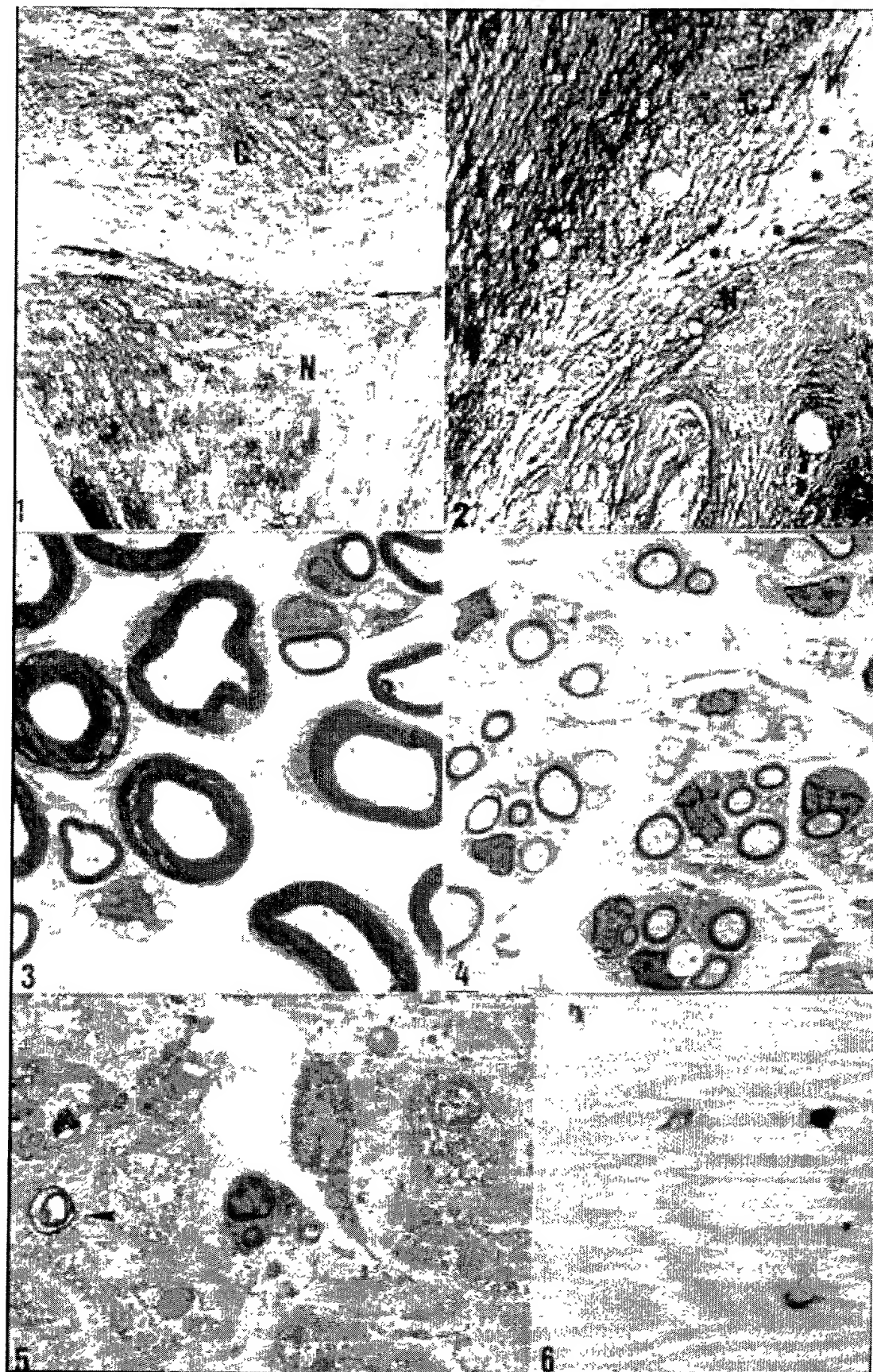


Fig. 14

Panel A: Body Weights of Normal Homozygous Mice vs. Wobblers Mice

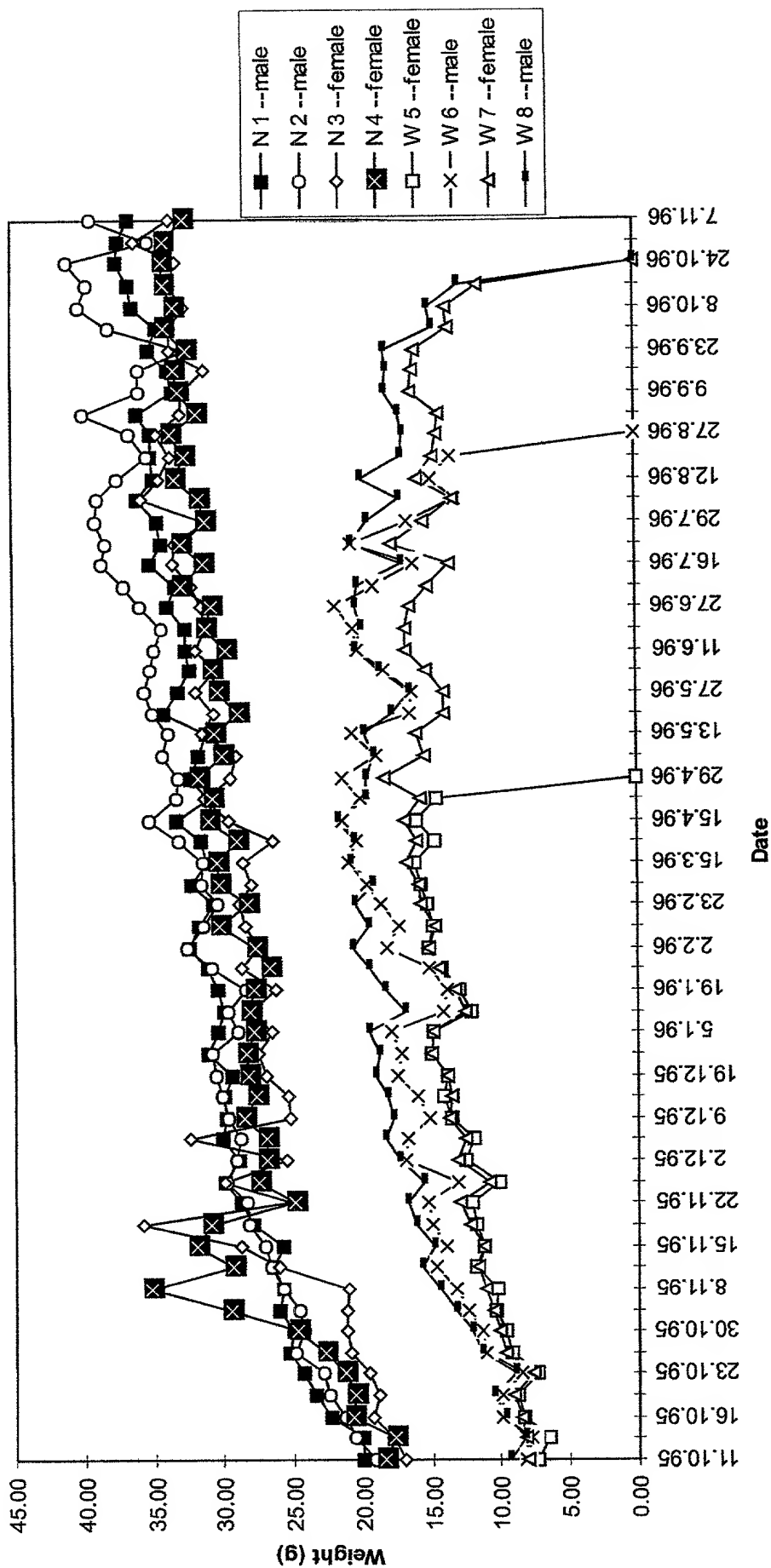


Fig. 15A

Panel B: Body Weights of Normal Heterozygous Mice vs. Wobblers Mice

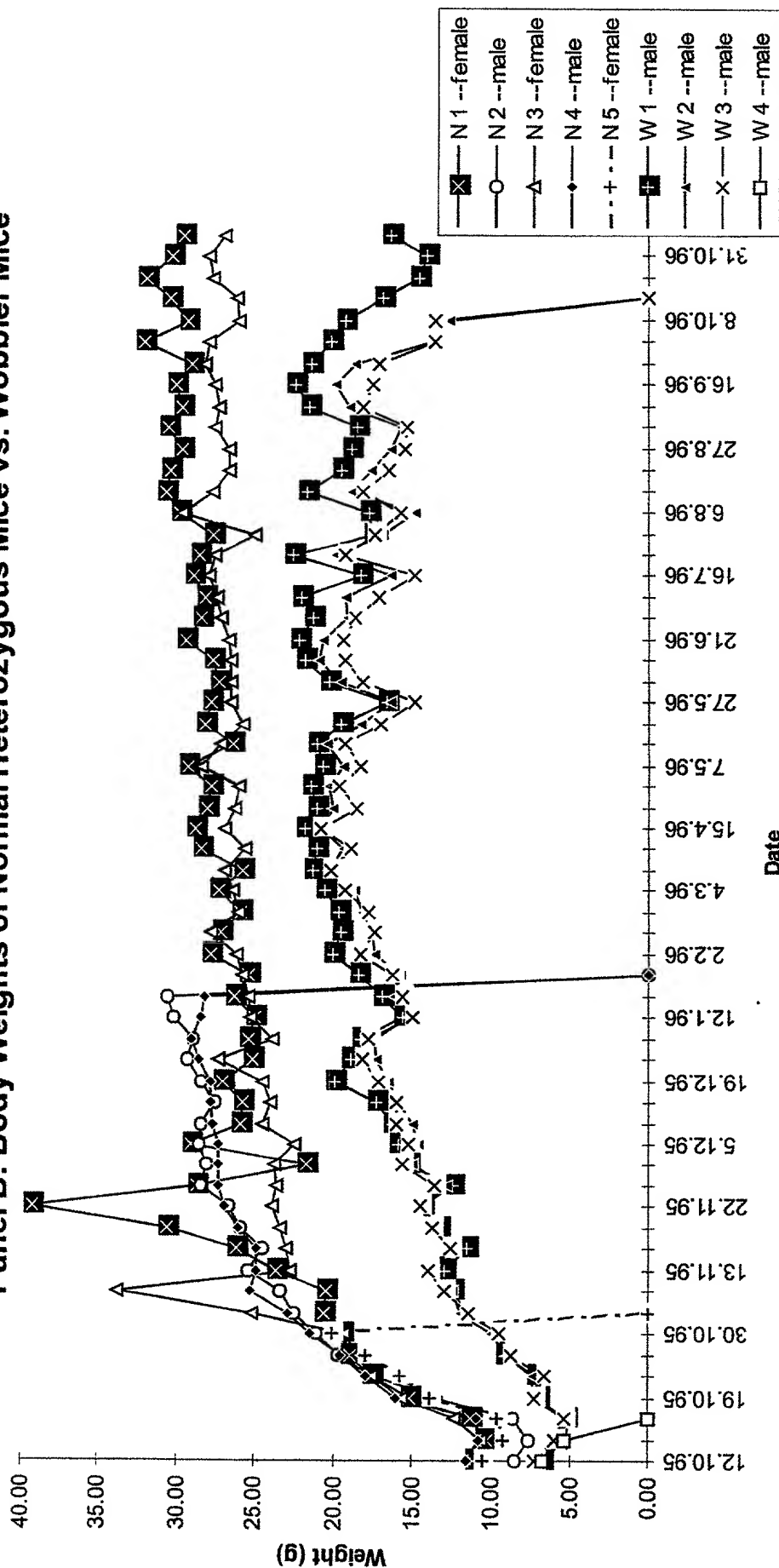


Fig. 15B

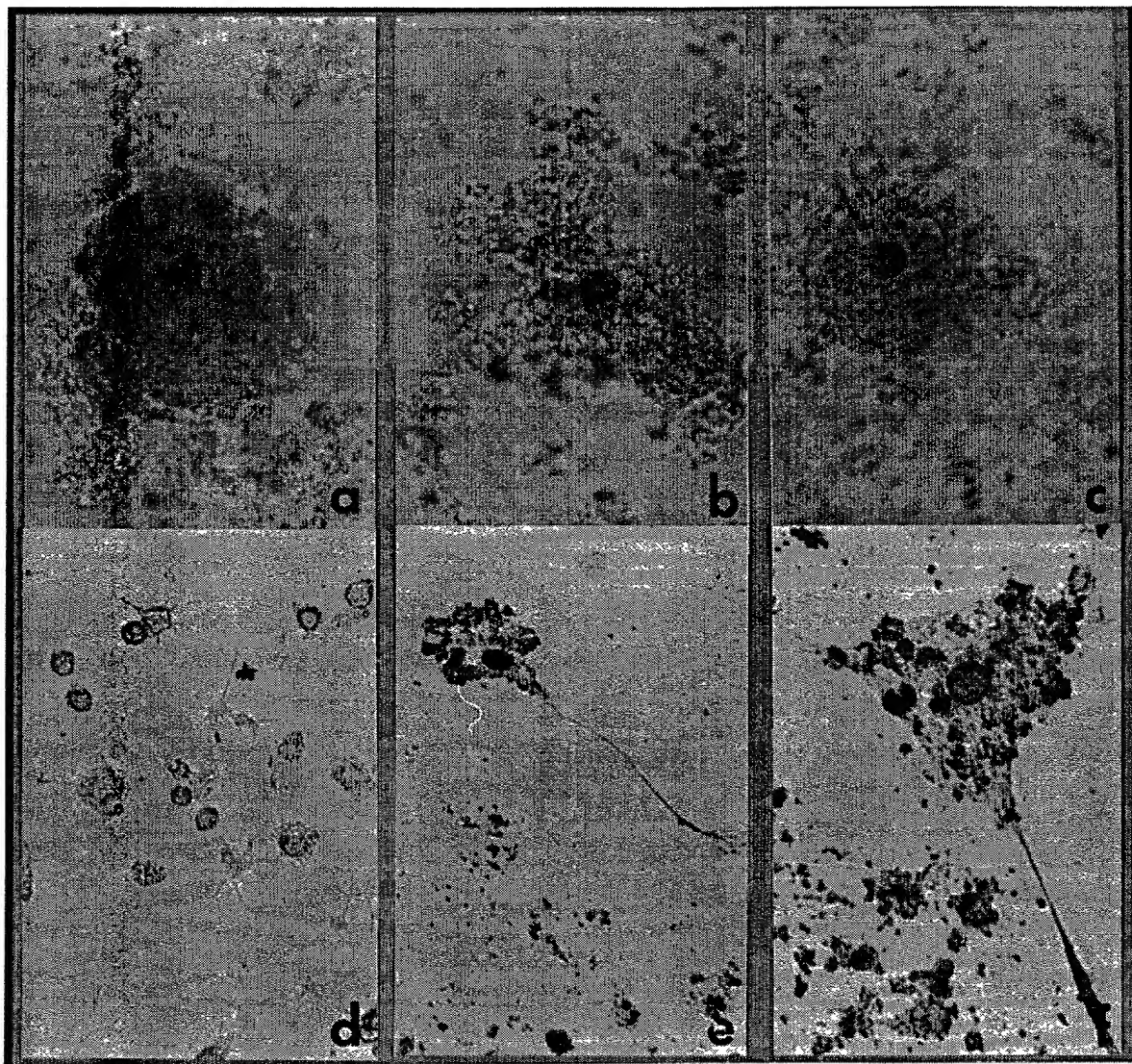


Fig. 16

6 -ve Control

3

F6 Cleared Protein

Fusion Protein

F6 Fusion Protein

F6 Class F3 Cleared Protein

Fig. 17

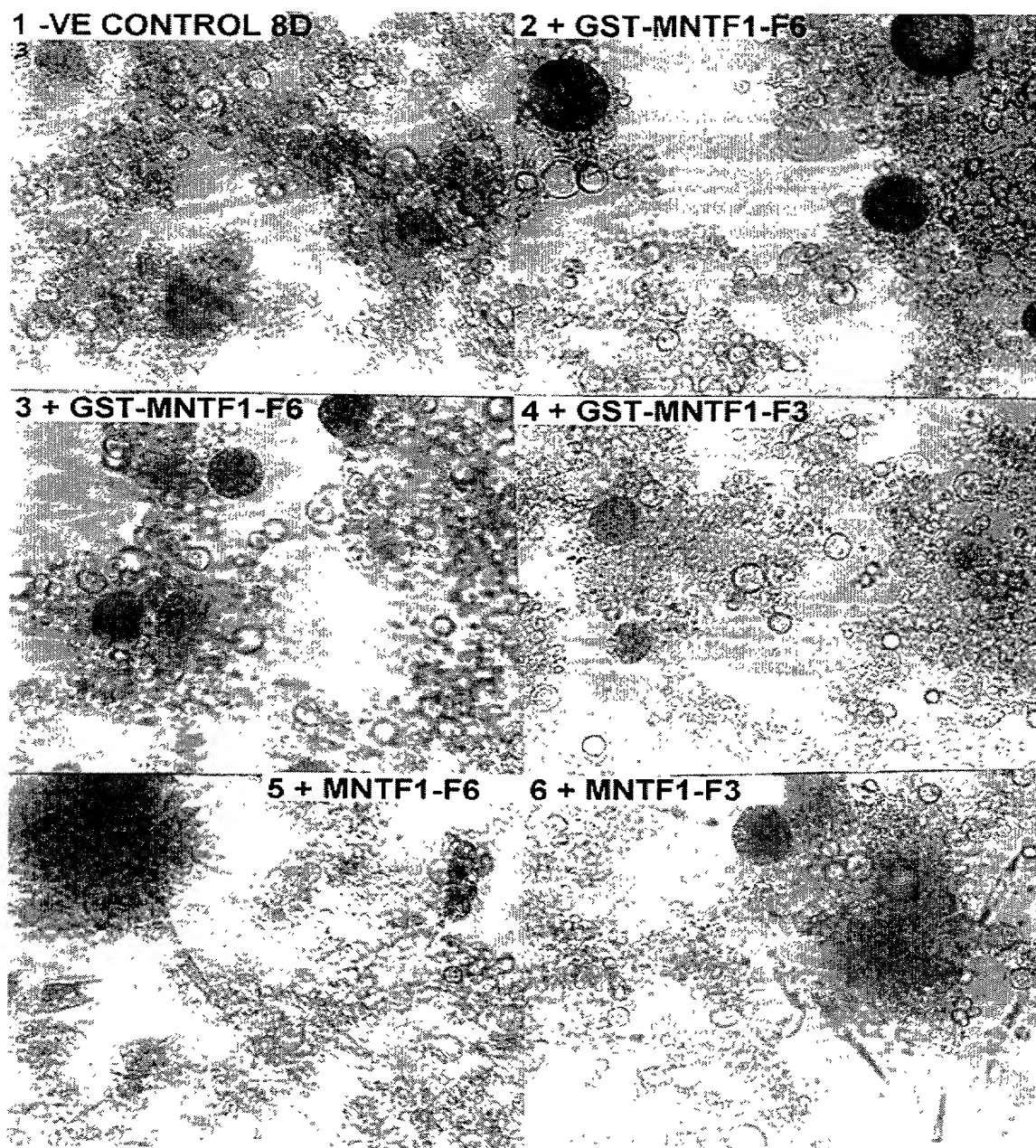


Fig. 17

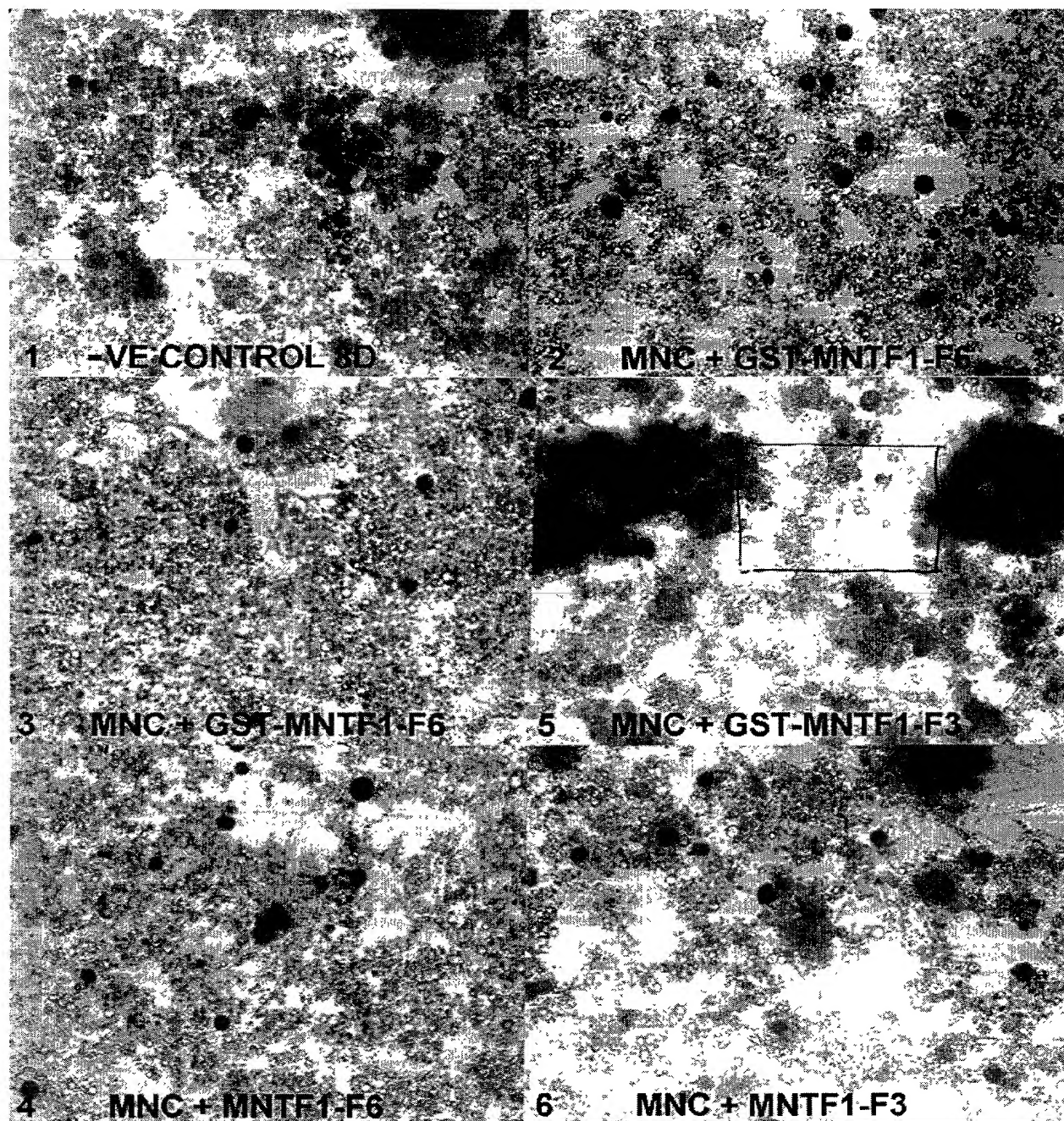
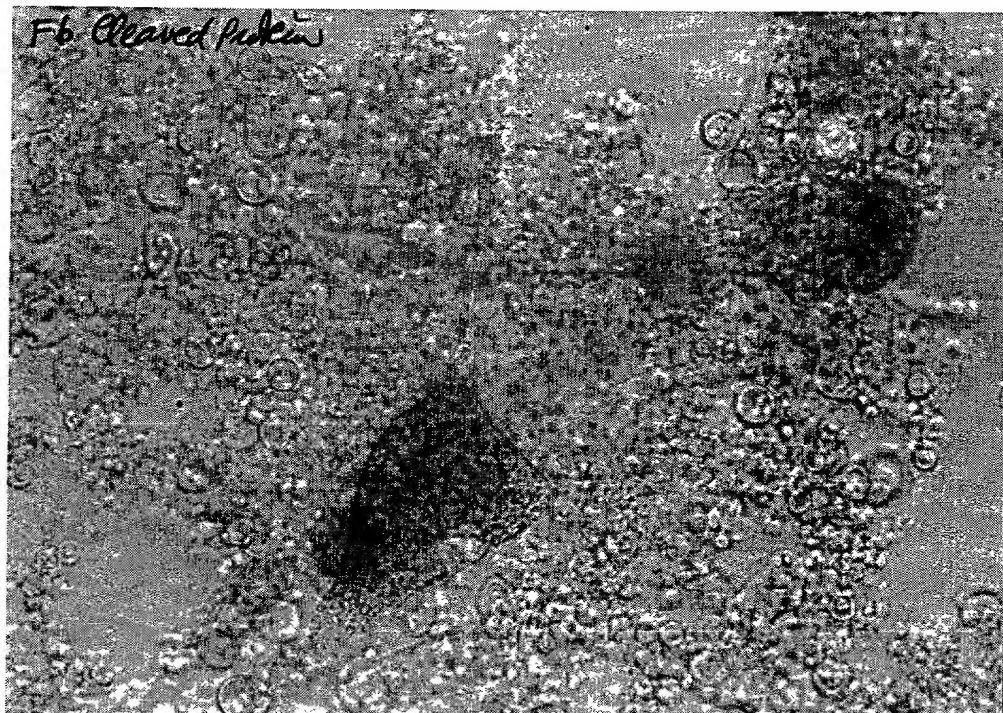
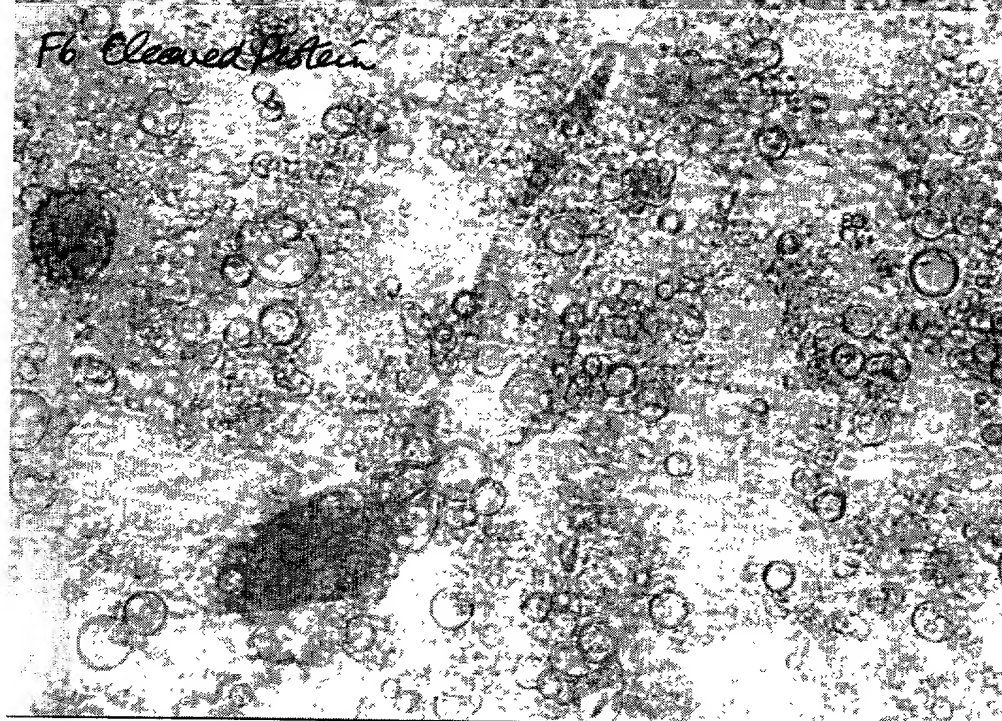


Fig. 18



PANEL 1



PANEL 2

Fig. 19

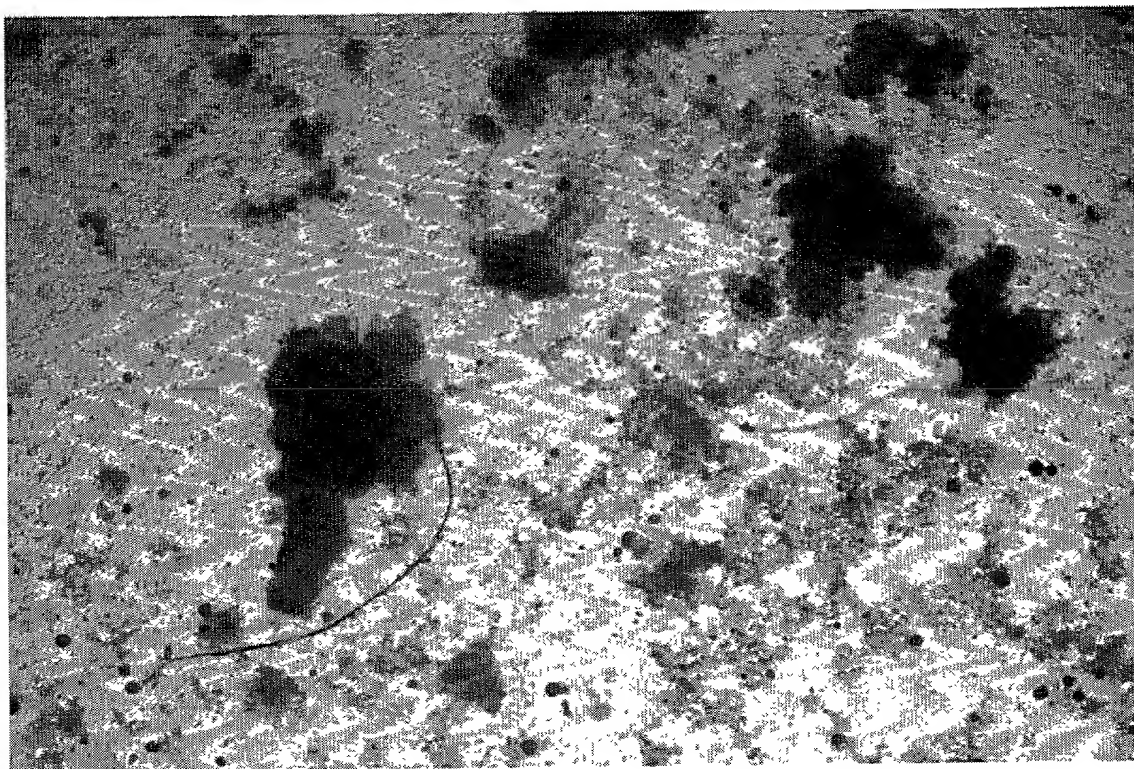


Fig. 20A

Fig. 20B

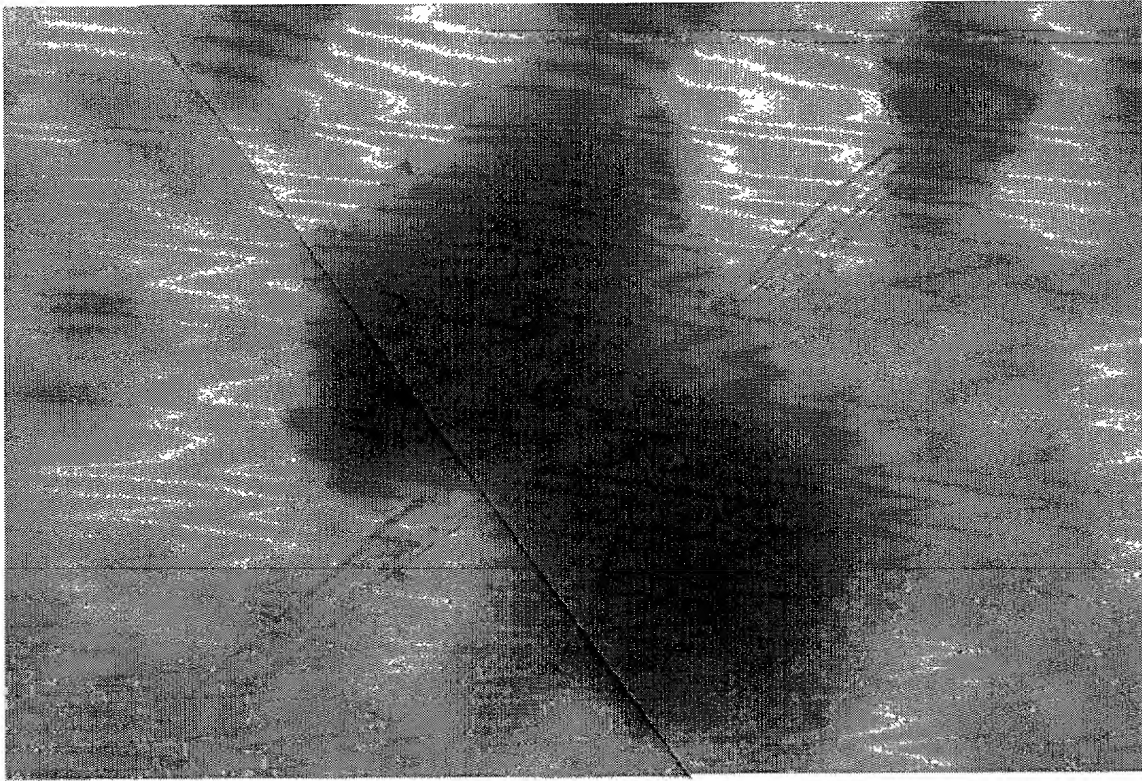


Fig. 20C

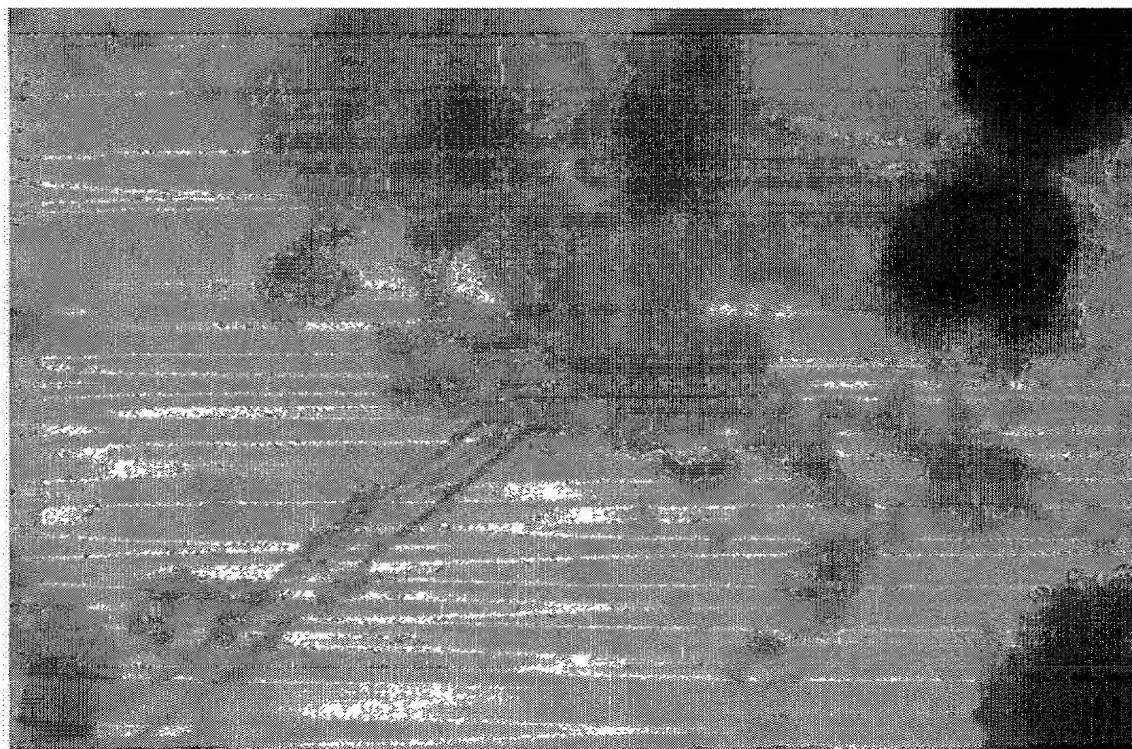


Fig. 20D